

WHAT IS CLAIMED IS:

*Sub B1* 1 An isolated nucleic acid encoding a tumor suppressor polypeptide  
2 p33ING2, wherein the polypeptide has greater than 70% amino acid sequence identity to  
3 a polypeptide comprising an amino acid sequence of SEQ ID NO:1.

1 (2) The isolated nucleic acid of claim 1, wherein the polypeptide  
2 selectively binds to polyclonal antibodies generated against a polypeptide comprising an  
3 amino acid sequence of SEQ ID NO:1.

1 (3) The isolated nucleic acid of claim 1, wherein the nucleic acid  
2 encodes a polypeptide comprising an amino acid sequence of SEQ ID NO:1.

1 (4) The isolated nucleic acid sequence of claim 1, wherein the nucleic  
2 acid comprises a nucleotide sequence of SEQ ID NO:2.

1 (5) The isolated nucleic acid of claim 1, wherein the nucleic acid is  
2 from a human.

*Sub B2* 1 (6) The isolated nucleic acid of claim 1, wherein the nucleic acid is  
2 amplified by primers that selectively hybridize under stringent hybridization conditions to  
3 the same sequence as degenerate primer sets encoding amino acid sequences selected  
4 from the group consisting of: SEQ ID NO:3 (MLGQQQQ) and SEQ ID NO:4  
5 (KKDRRSR).

1 (7) The isolated nucleic acid of claim 1, wherein the nucleic acid  
2 encodes a polypeptide having a molecular weight of about 28 kDa to about 38 kDa.

1 (8) An isolated nucleic acid encoding a tumor suppressor polypeptide  
2 p33ING2 that specifically hybridizes under stringent conditions to a nucleic acid  
3 comprising a nucleotide sequence of SEQ ID NO:2.

*Sub B3* 1 (9) The isolated nucleic acid of claim 1, wherein said nucleic acid  
2 selectively hybridizes under moderately stringent hybridization conditions to a nucleic  
3 acid comprising a nucleotide sequence of SEQ ID NO:2.

1           10. An isolated tumor suppressor polypeptide p33ING2, wherein the  
2 polypeptide has greater than 70 % amino acid sequence identity to a polypeptide  
3 comprising an amino acid sequence of SEQ ID NO:1.

1           11. The isolated tumor suppressor polypeptide of claim 10, wherein the  
2 polypeptide selectively binds to polyclonal antibodies generated against a polypeptide  
3 comprising an amino acid sequence of SEQ ID NO:1.

1           12. The isolated tumor suppressor polypeptide of claim 10, wherein the  
2 polypeptide comprises an amino acid sequence of SEQ ID NO:1.

1           13. The isolated tumor suppressor polypeptide of claim 10, wherein the  
2 polypeptide is from a human.

1           14. The isolated tumor suppressor polypeptide of claim 10, wherein the  
2 polypeptide is wild type p33ING2.  
*(D-130 later)*

1           15. An antibody that selectively binds to a p33ING2 polypeptide  
2 comprising an amino acid sequence of SEQ ID NO:1, but does not bind to a p33ING1  
3 polypeptide comprising an amino acid sequence of SEQ ID NO:8.

1           16. The antibody of claim 15, wherein the antibody is polyclonal.

1           17. An antibody that selectively binds to a p33ING1 polypeptide  
2 comprising an amino acid sequence of SEQ ID NO:8, but does not bind to a p33ING2  
3 polypeptide comprising an amino acid sequence of SEQ ID NO:1.

1           18. The antibody of claim 17, wherein the antibody is polyclonal.

1           19. The antibody of claim 15, wherein the antibody selectively binds to  
2 a p33ING2 polypeptide comprising the amino acid sequence of SEQ ID NO:5, but does  
3 not bind to a p33ING1 polypeptide comprising an amino acid sequence of SEQ ID NO:8.

1           20. An expression vector comprising the nucleic acid of claim 1.

1           21. A host cell transfected with the vector of claim 20.

1                   22. A method for identifying a compound that modulates a tumor  
2 suppressor polypeptide p33ING2, the method comprising the steps of:  
3                   (i) contacting the compound with a eukaryotic host cell or cell  
4 membrane in which has been expressed a tumor suppressor polypeptide p33ING2,  
5 wherein the polypeptide has greater than 70 % amino acid sequence identity to a  
6 polypeptide comprising an amino acid sequence of SEQ ID NO:1; and  
7                   (ii) determining the functional effect of the compound upon the  
8 cell or cell membrane expressing the polypeptide.

1                   23. The method of claim 22, wherein the polypeptide selectively binds  
2 to polyclonal antibodies generated against a polypeptide comprising an amino acid  
3 sequence of SEQ ID NO:1.

1                   24. The method of claim 22, wherein functional effect is determined by  
2 measuring changes in cell growth.

1                   25. The method of claim 22, wherein the polypeptide is recombinant.

1                   26. The method of claim 22, wherein the polypeptide is from a human.

1                   27. The method of claim 22, wherein the polypeptide comprises an  
2 amino acid sequence of SEQ ID NO:1.

1                   28. The method of claim 22, wherein the cell is an HCT116 human  
2 colon cancer cell line.

1                   29. The method of claim 22, wherein the cell has the missense  
2 p33ING2 sequence of a polypeptide comprising an amino acid sequence of SEQ ID  
3 NO:6.

1                   30. A method of inhibiting cellular proliferation, the method  
2 comprising  
3                   transducing a cell with an expression vector, the vector comprising a  
4 nucleic acid encoding a tumor suppressor polypeptide p33ING2, wherein the polypeptide  
5 has greater than 70 % amino acid sequence identity to a polypeptide comprising an amino  
6 acid sequence of SEQ ID NO:1. *and?*

*spans*

1           31. The method of claim 30, wherein the polypeptide selectively binds  
2 to polyclonal antibodies generated against a polypeptide comprising an amino acid  
3 sequence of SEQ ID NO:1.

1           32. The method of claim 30, wherein the nucleic acid encodes a  
2 polypeptide comprising an amino acid sequence of SEQ ID NO:1.

1           33. The method of claim 30, wherein the nucleic acid comprises a  
2 nucleotide sequence of SEQ ID NO:2.

1           34. The method of claim 30, wherein the nucleic acid is from a human.

1           35. The method of claim 30, wherein the nucleic acid encodes a  
2 polypeptide having a molecular weight of about 28 kDa to about 38 kDa.

1           36. The method of claim 30, wherein the cell is a HCT116 human  
2 colon cancer cell.

1           37. The method of claim 30, wherein the cell has a missense or null  
2 endogenous p33ING2 phenotype.

1           38. The method of claim 30, wherein the cell has a missense p33ING2  
2 sequence of a polypeptide comprising an amino acid sequence of SEQ ID NO:6.

1           39. A method of detecting the presence or absence of p33ING2 in  
2 mammalian tissue, the method comprising the steps of:

- 3                 (i) isolating a biological sample;
- 4                 (ii) contacting the biological sample with a p33ING2-specific  
5 reagent that selectively associates with p33ING2; and
- 6                 (iii) detecting the level of p33ING2-specific reagent that  
7 selectively associates with the sample.

1           40. The method of claim 39, wherein the p33ING2-specific reagent is  
2 selected from the group consisting of a p33ING2-specific antibody, a p33ING2-specific  
3 primer, and a p33ING2-specific nucleic acid probe.

1                  41. The method of claim 40, wherein the p33ING2-specific nucleic  
2 acid probe binds to a nucleic acid comprising a nucleotide sequence of SEQ ID NO:7, or  
3 to a nucleic acid comprising a nucleotide sequence of SEQ ID NO:2, or to a nucleic acid  
4 comprising a nucleotide sequence of SEQ ID NO:10.

1                  42. The method of claim 39, wherein the biological sample comprises  
2 intact chromosome 4q35.

1                  43. The method of claim 39, wherein the p33ING2-specific reagent  
2 detects nucleic acid.

1                  44. The method of claim 43, wherein the nucleic acid is a polymorphic  
2 variant of p33ING2.

1                  45. The method of claim 43, wherein the nucleic acid is RNA.

1                  46. The method of claim 39, wherein the p33ING2-specific reagent is  
2 an antibody that selectively binds to p33ING2.

1                  47. The method of claim 46, wherein the antibody is polyclonal.

1                  48. The method of claim 46, wherein the antibody selectively binds to  
2 a p33ING2 polypeptide comprising an amino acid sequence of SEQ ID NO:1, but not to a  
3 p33ING1 polypeptide comprising an amino acid sequence of SEQ ID NO:8.

1                  49. The antibody of claim 46, wherein the antibody selectively binds to  
2 a p33ING2 polypeptide comprising an amino acid sequence of SEQ ID NO:5, but does  
3 not bind to a p33ING1 polypeptide comprising an amino acid sequence of SEQ ID NO:8.

1                  50. A method of determining a test amount of p33ING2 in mammalian  
2 tissue, the method comprising the steps of:

- 3                  (i) isolating a biological sample;
- 4                  (ii) contacting the biological sample with a p33ING2-specific reagent
- 5                  that selectively associates with p33ING2; and
- 6                  (iii) comparing the test amount to a control.

1               51. The method of claim 50, wherein the control is an amount of  
2 p33ING2 in a normal cell.

1               52. The method of claim 50, wherein the p33ING2-specific reagent is  
2 selected from the group consisting of p33ING2-specific antibody, a p33ING2-specific  
3 primer; and p33ING2-specific nucleic acid probe.

1               53. A method of detecting the presence or absence of p33ING1 in  
2 mammalian tissue, the method comprising the steps of:

3                     (i) isolating a biological sample;  
4                     (ii) contacting the biological sample with a p33ING1-specific  
5 antibody that selectively binds to p33ING1 but not to p33ING2; and  
6                     (iii) detecting the level of p33ING1-specific antibody that  
7 selectively associates with the sample.

1               54. The method of claim 53, wherein the p33ING1-specific antibody is  
2 polyclonal.

1               55. A method of determining a test amount of p33ING1 in mammalian  
2 tissue, the method comprising the steps of:

3                     (i) isolating a biological sample;  
4                     (ii) contacting the biological sample with a p33ING1-specific antibody  
5 that selectively associates with p33ING1 but not to p33ING2; and  
6                     (iii) comparing the test amount to a control.

1               56. The method of claim 55, wherein the control is an amount of  
2 p33ING1 in a normal cell.

1               57. The method of claim 55, wherein the p33ING1-specific antibody is  
2 polyclonal.